

## SEARCH REQUEST FORM

6-114

FONDA 6-30-99

Examiner # (Mandatory): \_\_\_\_\_ Requester's Full Name: \_\_\_\_\_

Art Unit \_\_\_\_\_ Location (Bldg/Room#): 7D03 Phone (circle 305 306 308) \_\_\_\_\_Serial Number: 19/300173 Results Format Preferred (circle): PAPER DISK E-MAIL

Title of Invention \_\_\_\_\_

Inventors (please provide full names): \_\_\_\_\_  
\_\_\_\_\_

Earliest Priority Date: \_\_\_\_\_

Keywords (include any known synonyms registry numbers, explanation of initialisms):

see claim 2 for polysaccharides.

see claims 5, 6, and 8 for hydrophobic agents.

## Search Topic:

Please write detailed statement of the search topic, and the concept of the invention. Describe as specifically as possible the subject matter to be searched. Define any terms that may have a special meaning. Give examples of relevant citations, authors, etc., if known. You may include a copy of the abstract and the broadcast or most relevant claim(s).

Please search attached claims 1-20.

Note that the method claims are limited to heparin as the polysaccharide.

## STAFF USE ONLY

Searcher: \_\_\_\_\_

## Type of Search

## Vendors (include cost where applicable)

Searcher Phone #: -49-58 N.A. Sequence STN

Searcher Location: \_\_\_\_\_

 A.A. Sequence Questel/Orbit

Date Picked Up: \_\_\_\_\_

 Structure (#) Lexis/NexisDate Completed: 7/1/ Bibliographic WWW/InternetClerical Prep Time: 17:15 Litigation In-house sequence systems (list)Terminal Time: 85:150 Fulltext DialogNumber of Databases: 2 Procurement Dr. Link Other Westlaw Other (specify)

=> fil reg

FILE 'REGISTRY' ENTERED AT 12:24:33 ON 06 JUL 1999  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
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STRUCTURE FILE UPDATES: 03 JUL 99 HIGHEST RN 227003-54-5  
DICTIONARY FILE UPDATES: 06 JUL 99 HIGHEST RN 227003-54-5

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 13, 1999

Please note that search-term pricing does apply when  
conducting SmartSELECT searches.

=> d his

(FILE 'HOME' ENTERED AT 11:08:43 ON 06 JUL 1999)

SET COST OFF  
SET AUHELP OFF

FILE 'HCAPLUS' ENTERED AT 11:08:52 ON 06 JUL 1999

E BYUN Y/AU  
L1 14 S E3,E30  
L2 10 S L1 AND HEPARIN  
E LEE Y/AU  
L3 293 S E3,E27  
E LEE YONG/AU  
L4 40 S E3,E83  
E LEE YONGKYU/AU  
L5 1 S E3  
L6 1 S L3-L5 AND HEPARIN  
L7. 1 S L1 AND L3-L5  
L8 1 S L2,L6 AND L7  
L9 0 S L1 AND P/DT  
L10 0 S L3-L5 AND P/DT  
E KR98/AP, PRN  
E KR98-19469/AP, PRN

FILE 'INPADOC' ENTERED AT 11:14:58 ON 06 JUL 1999

E KR98-19469/AP, PRN

FILE 'WPIDS' ENTERED AT 11:15:14 ON 06 JUL 1999

E KR98-19469/AP, PRN

FILE 'HCAPLUS' ENTERED AT 11:15:31 ON 06 JUL 1999

L11 243 S SODIUM HEPARIN  
L12 196202 S CELLULOSE  
L13 171 S HYDROXYMETHYLCELLULOSE  
L14 1258 S HYDROXYPROPYLCELLULOSE  
L15 4220 S CHOLIC ACID  
L16 2476 S DEOXYCHOLIC ACID  
L17 2494 S CHENODEOXYCHOLIC ACID  
L18 1428 S LITHOCHOLIC ACID  
L19 77 S URSOCHOLIC ACID  
L20 1589 S URSODEOXYCHOLIC ACID  
L21 12 S ISOURSODEOXYCHOLIC ACID  
L22 3 S LAGODEOXYCHOLIC ACID

L23        640 S GLYCOCHOLIC ACID  
L24        1465 S TAUROCHOLIC ACID  
L25        277 S GLYCODEOXYCHOLIC ACID  
L26        296 S GLYCOCHENODEOXYCHOLIC ACID  
L27        302 S DEHYDROCHOLIC ACID  
L28        134 S HYOCHOLIC ACID  
L29        383 S HYODEOXYCHOLIC ACID  
L30        1051 S CHOLESTANOL  
L31        452 S COPROSTANOL  
L32        90458 S CHOLESTEROL  
L33        98 S EPICHOLESTEROL  
L34        3276 S ERGOSTEROL  
L35        471 S ERGOCALCIFEROL  
L36        12138 S BUTYRIC ACID  
L37        3211 S VALERIC ACID  
L38        2586 S CAPROIC ACID  
L39        2331 S CAPRYLIC ACID  
L40        1954 S CAPRIC ACID  
L41        6952 S LAURIC ACID  
L42        5498 S MYRISTIC ACID  
L43        15475 S PALMITIC ACID  
L44        33137 S STEARIC ACID  
L45        12561 S POLYETHYLENEOXIDE OR POLY ETHYLENEOXIDE OR POLYETHYLENE OXIDE  
L46        1480 S POLYEPSIOLONCAPROLACTONE OR POLYEPSILON CAPROLACTONE OR POLY  
L47        156 S CAPROLACTONE (L) (ETHYLENEOXIDE OR ETHYLENE OXIDE)  
L48        20334 S ETHYLENE VINYL ACETATE

FILE 'REGISTRY' ENTERED AT 11:32:44 ON 06 JUL 1999  
L49        5 S 9005-49-6 OR 9041-08-1 OR 9004-34-6 OR 37353-59-6 OR 9004-64-

FILE 'HCAPLUS' ENTERED AT 11:33:21 ON 06 JUL 1999  
L50        68515 S L49  
L51        225243 S L11-L14 OR HEPARIN OR (SULFONAT? OR SULPHONAT?) (L) ?SACCHARIDE

FILE 'REGISTRY' ENTERED AT 11:35:34 ON 06 JUL 1999  
L52        15 S 81-25-4 OR 83-44-3 OR 474-25-9 OR 434-13-9 OR 2955-27-3 OR 12

FILE 'HCAPLUS' ENTERED AT 11:38:06 ON 06 JUL 1999  
L53        423 S (L52 OR L15-L29 OR BILE ACID) AND L51

FILE 'REGISTRY' ENTERED AT 11:39:20 ON 06 JUL 1999  
L54        6 S 80-97-7 OR 360-68-9 OR 57-88-5 OR 474-77-1 OR 57-87-4 OR 50-1

FILE 'HCAPLUS' ENTERED AT 11:39:30 ON 06 JUL 1999  
L55        2486 S (L54 OR L30-L35 OR STEROL) AND L51

FILE 'REGISTRY' ENTERED AT 11:40:35 ON 06 JUL 1999

FILE 'REGISTRY' ENTERED AT 11:40:50 ON 06 JUL 1999  
L56        9 S 107-92-6 OR 109-52-4 OR 142-62-1 OR 124-07-2 OR 143-07-7 OR 5

FILE 'HCAPLUS' ENTERED AT 11:41:02 ON 06 JUL 1999  
L57        2648 S (L56 OR L36-L44 OR ALKANOIC ACID) AND L51  
L58        685 S POLYSACCHARIDE AND HYDROPHOBIC  
L59        48 S L58 AND COVALEN?  
L60        5250 S L53, L55, L57  
L61        48 S L60 AND COVALEN?  
L62        91 S L59, L61  
L63        25 S L62 AND PHARMACEUT?/SC, SX, CW, BI, AB

L64 5 S L62 AND L52  
L65 19 S L62 AND L54  
L66 14 S L62 AND L56  
L67 34 S L64-L66  
L68 24 S L67 AND L50  
L69 24 S L68 AND COVALEN?

FILE 'HCAPLUS' ENTERED AT 11:51:45 ON 06 JUL 1999

FILE 'REGISTRY' ENTERED AT 11:52:26 ON 06 JUL 1999

L70 1 S 107596-21-4  
L71 127 S 75-21-8/CRN AND 502-44-3/CRN  
L72 3 S L71 AND 2/NC  
E POLYURETHANE/CN  
E URETHANE/CN  
E ETHYLENE VINYL ACETATE/CN  
E ETHYLENE VINYLACETATE/CN  
E ETHYLENEVINYLACETATE/CN  
L73 1 S 30174-06-2  
L74 1 S 24937-78-8  
L75 1760 S 74-85-1/CRN AND 108-05-4/CRN  
L76 6 S L75 AND 2/NC  
L77 3 S L76 NOT MAN/CI  
L78 6 S L72,L77  
L79 6 S L73,L74,L78

FILE 'HCAPLUS' ENTERED AT 11:58:36 ON 06 JUL 1999

L81 245673 S L\*\*\*,L45-L48, POLYURETHANE OR URETHANE OR SILICONE OR SILOXANE  
L82 26057 S L79  
L83 268821 S L81,L82  
L84 460 S L83 AND L60  
L85 3 S L84 AND COVALEN?

FILE 'REGISTRY' ENTERED AT 12:08:44 ON 06 JUL 1999

L86 2 S 9005-49-6 OR 9041-08-1

FILE 'HCAPLUS' ENTERED AT 12:09:08 ON 06 JUL 1999

L87 15840 S L86  
L88 505 S L87 AND L53,L55,L57,L58  
L89 13 S L88 AND COVALEN?  
L90 5 S L89 AND 63/SC,SX  
L91 8 S L89 NOT L90  
L92 374 S L87 AND L52,L54,L56  
L93 0 S L92 AND L82  
L94 11 S L92 AND (SUSTAIN? OR CONTROL?) (L)RELEASE?  
L95 4 S L94 AND (PERCUTANEOUS OR HDL OR LIPOLYTIC OR STREPTOZOTOCIN) /  
L96 7 S L94 NOT L95  
L97 11 S L90,L96  
L98 33 S L87 AND L82  
L99 5 S L98 AND COVALEN?  
L100 4 S L99 AND 63/SC,SX  
L101 15 S L97,L100  
L102 29 S L98 NOT L101  
L103 28 S L102 AND 63/SC,SX  
L104 0 S L103 AND L60  
SEL HIT RN L101

FILE 'REGISTRY' ENTERED AT 12:23:43 ON 06 JUL 1999

L105 10 S E1-E10

FILE 'REGISTRY' ENTERED AT 12:24:33 ON 06 JUL 1999

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L105 ANSWER 1 OF 10 REGISTRY COPYRIGHT 1999 ACS  
RN 24937-78-8 REGISTRY  
CN Acetic acid ethenyl ester, polymer with ethene (9CI) (CA INDEX NAME)  
OTHER CA INDEX NAMES:  
CN Acetic acid vinyl ester, polymer with ethylene (8CI)  
OTHER NAMES:  
CN 1010VN3  
CN 1900W  
CN 204CS95  
CN 3043H  
CN 3130SB  
CN 3135F  
CN 33G1A  
CN 3507C  
CN 547D  
CN 54C  
CN 84D  
CN A 400  
CN A 400 (vinyl polymer)  
CN A 416  
CN A 416 (polymer)  
CN A 443/31  
CN A 9918  
CN AC 400  
CN AC 400A  
CN AC 401  
CN AC 405  
CN AC 405S  
CN AC 405T  
CN AC 410  
CN AC 430  
CN AC 440  
CN AC-P 400  
CN AD 1790-15  
CN AD 37JD965  
CN AD 37P147  
CN AD 37P295  
CN AD 37P66  
CN AD 5  
CN AD 511  
CN Adcote 1790  
CN Adcote 295G  
CN Adcote 33-131  
CN Adcote 33G1A  
CN Adcote 37JD610  
CN Adcote 37R610  
CN Adcote AD 37P295  
CN Adcote AD 37P295J  
CN Adcote X 17  
CN Adcote X 37T77  
CN Adeva 629  
CN Admer AT 589  
CN Admer NE 100

CN AE 221

CN Aibon AE 20

ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for  
DISPLAYDR 158707-29-0, 53637-14-2, 58252-58-7, 58858-06-3, 123757-93-7, 60529-82-0,  
64296-24-8, 129291-57-2, 129496-11-3, 97445-08-4, 103470-05-9,  
103843-24-9, 51312-30-2, 141255-84-7, 111367-02-3, 112820-85-6,  
137802-65-4, 74566-07-7, 148709-22-2, 78355-65-4, 144046-56-0,  
144246-76-4, 81406-40-8, 86904-51-0, 88024-59-3, 39457-29-9, 116811-82-6,  
117217-63-7, 117217-64-8, 117217-68-2, 117313-46-9, 183815-97-6

MF (C4 H6 O2 . C2 H4)x

CI PMS, COM

PCT Polyolefin, Polyvinyl

LC STN Files: AGRICOLA, ANABSTR, ASMDATA\*, BIOBUSINESS, BIOSIS, CA,  
CANCERLIT, CAPLUS, CEN, CHEMCATS, CHEMLIST, CBNB, CIN, CSCHEM, DDFU,  
DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MSDS-OHS, PIRA,  
PLASPEC\*, PROMT, TOXLINE, TOXLIT, USPATFULL

(\*File contains numerically searchable property data)

Other Sources: DSL\*\*, TSCA\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

CM 1

CRN 108-05-4

CMF C4 H6 O2

AcO-CH=CH2

CM 2

CRN 74-85-1

CMF C2 H4

H2C=CH2

25899 REFERENCES IN FILE CA (1967 TO DATE)

2374 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

25983 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:25799

REFERENCE 2: 131:25720

REFERENCE 3: 131:23350

REFERENCE 4: 131:22889

REFERENCE 5: 131:21343

REFERENCE 6: 131:21225

REFERENCE 7: 131:20892

REFERENCE 8: 131:20359

REFERENCE 9: 131:20124

REFERENCE 10: 131:20123

L105 ANSWER 2 OF 10 REGISTRY COPYRIGHT 1999 ACS  
RN 9041-08-1 REGISTRY  
CN Heparin, sodium salt (8CI, 9CI) (CA INDEX NAME)

OTHER NAMES:

CN Alfa 87-120  
CN Alfa 87-163  
CN Alfa 87-198  
CN Alfa 87-81  
CN Alfa 88-247  
CN Ardeparin sodium  
CN Bemiparin sodium  
CN Dalteparin sodium  
CN Danaparoid sodium  
CN Depo-Heparin  
CN Enoxaparin sodium  
CN Fragmin  
CN Fragmin IV  
CN H 2149  
CN Hed-Heparin  
CN Heparin sodium  
CN Hepathrom  
CN Inno-Hep  
CN Kabi 2165  
CN LHN 1  
CN Liquaemin sodium  
CN Liquemin  
CN Logiparin  
CN Lovenox  
CN Minolteparin sodium  
CN Normiflo  
CN Parnaparin sodium  
CN PK 10169  
CN Pularin  
CN Reviparin sodium  
CN RO 11  
CN RP 54563  
CN Sodium acid heparin  
CN Sodium heparin  
CN Sodium heparinate  
CN Tinzaparin sodium  
CN WY 90493RD

DR 12656-11-0, 101921-26-0, 102785-31-9

MF Unspecified

CI PMS, COM, MAN

PCT Manual registration, Polyester, Polyester formed

LC STN Files: ADISINSIGHT, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, CA,  
CABA, CAPLUS, CHEMCATS, CHEMLIST, CBNB, CIN, CSCHEM, DDFU, DRUGNL,  
DRUGPAT, DRUGU, DRUGUPDATES, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA,  
MEDLINE, MRCK\*, MSDS-OHS, NIOSHTIC, PHAR, PROMT, RTECS\*, TOXLINE,  
TOXLIT, USAN, USPATFULL

(\*File contains numerically searchable property data)

Other Sources: DSL\*\*, TSCA\*\*, WHO

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

789 REFERENCES IN FILE CA (1967 TO DATE)

67 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

792 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:23343

REFERENCE 2: 130:351211

REFERENCE 3: 130:332537

REFERENCE 4: 130:332211

REFERENCE 5: 130:324355

REFERENCE 6: 130:320843

REFERENCE 7: 130:242240

REFERENCE 8: 130:227804

REFERENCE 9: 130:227796

REFERENCE 10: 130:217977

L105 ANSWER 3 OF 10 REGISTRY COPYRIGHT 1999 ACS

RN 9005-49-6 REGISTRY

CN Heparin (8CI, 9CI) (CA INDEX NAME)

OTHER NAMES:

CN .alpha.-Heparin

CN Bemiparin

CN Certoparin

CN Clexane

CN Clivarin

CN Clivarine

CN CY 216

CN CY 222

CN Dalteparin

CN Enoxaparin

CN Fluxum

CN FR 860

CN Fragmin A

CN Fragmin B

CN Fraxiparin

CN Heparin sulfate

CN Heparinic acid

CN KB 101

CN Novoheparin

CN OP 386

CN OP 622

CN Pabyrn

CN Parnaparin

CN Parvoparin

CN Reviparin

CN Sandoparin

CN Sublingula

CN Vetren

CN Vitrum AB

DR 9075-96-1, 11078-24-3, 11129-39-8, 104521-37-1, 37324-73-5, 91449-79-5

MF Unspecified  
 CI PMS, COM, MAN  
 PCT Manual registration, Polyester, Polyester formed  
 LC STN Files: ADISINSIGHT, AGRICOLA, AIDSLINE, ANABSTR, BIOBUSINESS,  
     BIOSIS, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMLIST,  
     CBNB, CIN, CSCHEM, DDFU, DRUGNL, DRUGPAT, DRUGU, DRUGUPDATES, EMBASE,  
     HSDB\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, NAPRALERT,  
     NIOSHTIC, PIRA, PHAR, PROMT, RTECS\*, TOXLINE, TOXLIT, USAN, USPATFULL  
     (\*File contains numerically searchable property data)  
 Other Sources: DSL\*\*, EINECS\*\*, WHO  
     (\*\*Enter CHEMLIST File for up-to-date regulatory information)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*  
 15277 REFERENCES IN FILE CA (1967 TO DATE)  
 1542 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 15300 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:23585

REFERENCE 2: 131:23577

REFERENCE 3: 131:23532

REFERENCE 4: 131:23508

REFERENCE 5: 131:23483

REFERENCE 6: 131:19223

REFERENCE 7: 131:19222

REFERENCE 8: 131:17420

REFERENCE 9: 131:16976

REFERENCE 10: 131:16875

L105 ANSWER 4 OF 10 REGISTRY COPYRIGHT 1999 ACS

RN 2955-27-3 REGISTRY

CN Cholan-24-oic acid, 3,7,12-trihydroxy-, (3.alpha.,5.beta.,7.beta.,12.alpha.)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5.beta.-Cholan-24-oic acid, 3.alpha.,7.beta.,12.alpha.-trihydroxy- (8CI)

CN 5.beta.-Cholanic acid, 3.alpha.,7.beta.,12.alpha.-trihydroxy- (7CI)

OTHER NAMES:

CN 3.alpha.,7.beta.,12.alpha.-Trihydroxy-5.beta.-cholanic acid

CN 3.alpha.,7.beta.,12.alpha.-Trihydroxy-5.beta.-cholanoic acid

CN 3.alpha.,7.beta.,12.alpha.-Trihydroxycholanic acid

CN 7-Epicholic acid

CN 7.beta.-Hydroxyisocholic acid

CN Ursocholic acid

FS STEREOSEARCH

DR 121571-08-2

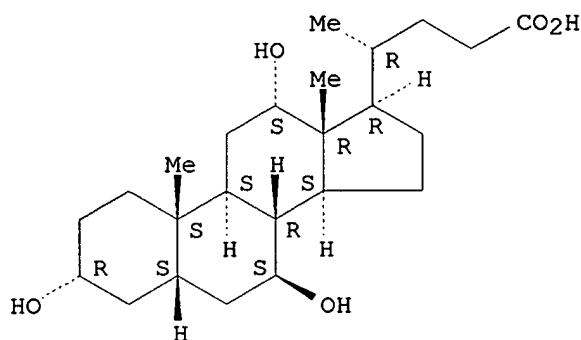
MF C24 H40 O5

CI COM

LC STN Files: BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CAOLD, CAPLUS, CASREACT, CHEMINFORMRX, DETHERM\*, DDFU, DRUGU, EMBASE, GMELIN\*, IPA, MEDLINE, NAPRALERT, TOXLINE, TOXLIT, USPATFULL

(\*File contains numerically searchable property data)

Absolute stereochemistry.



154 REFERENCES IN FILE CA (1967 TO DATE)  
 6 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 154 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
 7 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 130:308722

REFERENCE 2: 130:301714

REFERENCE 3: 129:250255

REFERENCE 4: 129:245326

REFERENCE 5: 128:326328

REFERENCE 6: 128:280428

REFERENCE 7: 128:215953

REFERENCE 8: 128:126586

REFERENCE 9: 128:39642

REFERENCE 10: 128:21062

L105 ANSWER 5 OF 10 REGISTRY COPYRIGHT 1999 ACS

RN 544-63-8 REGISTRY

CN Tetradecanoic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Myristic acid (8CI)

OTHER NAMES:

CN 1-Tridecanecarboxylic acid

CN Edenor C 14

CN Emery 655

CN Hystrene 9014

CN Kortacid 1499

CN n-Tetradecan-1-oic acid

CN n-Tetradecanoic acid

CN n-Tetradecoic acid

CN NAA 104

CN NAA 142  
 CN Neo-Fat 14  
 CN Philacid 1400  
 CN Prifac 2942  
 CN Univol U 316S  
 FS 3D CONCORD  
 DR 45184-05-2  
 MF C14 H28 O2  
 CI COM  
 LC STN Files: AGRICOLA, AIDSLINE, ANABSTR, APILIT, APILIT2, APIPAT,  
     APIPAT2, BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CABAB, CANCERLIT, CAOLD,  
     CAPLUS, CASREACT, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM,  
     CSNB, DETHERM\*, DDFU, DIPPR\*, DRUGU, EMBASE, GMELIN\*, HODOC\*, HSDB\*,  
     IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, NAPRALERT,  
     NIOSHTIC, PDLCOM\*, PIRA, PROMT, RTECS\*, SPECINFO, TOXLINE, TOXLIT,  
     TRCTHERMO\*, TULSA, USPATFULL, VTB  
     (\*File contains numerically searchable property data)  
 Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\*  
     (\*\*Enter CHEMLIST File for up-to-date regulatory information)

HO2C—(CH2)12—Me

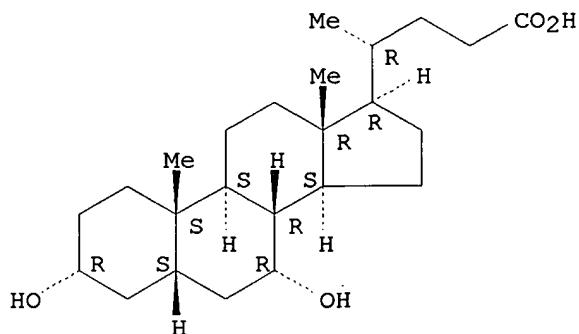
11215 REFERENCES IN FILE CA (1967 TO DATE)  
 483 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 11223 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
 13 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 131:23512  
 REFERENCE 2: 131:23217  
 REFERENCE 3: 131:22112  
 REFERENCE 4: 131:18402  
 REFERENCE 5: 131:18334  
 REFERENCE 6: 131:18183  
 REFERENCE 7: 131:18150  
 REFERENCE 8: 131:18135  
 REFERENCE 9: 131:18097  
 REFERENCE 10: 131:18094

L105 ANSWER 6 OF 10 REGISTRY COPYRIGHT 1999 ACS  
 RN 474-25-9 REGISTRY  
 CN Cholan-24-oic acid, 3,7-dihydroxy-, (3.alpha.,5.beta.,7.alpha.)- (9CI)  
     (CA INDEX NAME)  
 OTHER CA INDEX NAMES:  
 CN 5.beta.-Cholan-24-oic acid, 3.alpha.,7.alpha.-dihydroxy- (8CI)  
 OTHER NAMES:  
 CN 3.alpha.,7.alpha.-Dihydroxy-5.beta.-cholan-24-oic acid  
 CN 3.alpha.,7.alpha.-Dihydroxy-5.beta.-cholanic acid  
 CN 3.alpha.,7.alpha.-Dihydroxy-5.beta.-cholanoic acid

CN 3.alpha.,7.alpha.-Dihydroxycholanic acid  
 CN 7.alpha.-Hydroxylithocholic acid  
 CN Anthropodeoxycholic acid  
 CN Anthropodesoxycholic acid  
 CN Anthropododesoxycholic acid  
 CN CDC  
 CN Chendol  
 CN Chenic acid  
 CN Chenodeoxycholic acid  
 CN Chenodesoxycholic acid  
 CN Chenodiol  
 CN Gallodesoxycholic acid  
 FS STEREOSEARCH  
 MF C24 H40 O4  
 CI COM  
 LC STN Files: AGRICOLA, AIDSLINE, ANABSTR, BEILSTEIN\*, BIOBUSINESS, BIOSIS,  
     CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX,  
     CHEMLIST, CBNB, CIN, CSCHEM, CSNB, DETHERM\*, DDFU, DRUGU, EMBASE,  
     HODOC\*, HSDB\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS,  
     NAPRALERT, NIOSHTIC, PROMT, RTECS\*, SPECINFO, TOXLINE, TOXLIT, USAN,  
     USPATFULL  
     (\*File contains numerically searchable property data)  
 Other Sources: DSL\*\*, EINECS\*\*, WHO  
     (\*\*Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.



2964 REFERENCES IN FILE CA (1967 TO DATE)  
 143 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 2968 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
 20 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

- |           |    |            |
|-----------|----|------------|
| REFERENCE | 1: | 131:18391  |
| REFERENCE | 2: | 131:15856  |
| REFERENCE | 3: | 131:14194  |
| REFERENCE | 4: | 131:2721   |
| REFERENCE | 5: | 130:350237 |
| REFERENCE | 6: | 130:345049 |
| REFERENCE | 7: | 130:322477 |

REFERENCE 8: 130:309449

REFERENCE 9: 130:308722

REFERENCE 10: 130:306386

L105 ANSWER 7 OF 10 REGISTRY COPYRIGHT 1999 ACS

RN 128-13-2 REGISTRY

CN Cholan-24-oic acid, 3,7-dihydroxy-, (3.alpha.,5.beta.,7.beta.)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5.beta.-Cholan-24-oic acid, 3.alpha.,7.beta.-dihydroxy- (8CI)

OTHER NAMES:

CN 3.alpha.,7.beta.-Dihydroxy-5.beta.-cholan-24-oic acid

CN 3.alpha.,7.beta.-Dihydroxy-5.beta.-cholanic acid

CN 3.alpha.,7.beta.-Dihydroxy-5.beta.-cholanoic acid

CN 3.alpha.,7.beta.-Dihydroxycholanic acid

CN 7.beta.-Hydroxylithocholic acid

CN Actigall

CN Desocol

CN Deursil

CN Urso

CN Ursocholic acid, deoxy-

CN Ursodeoxycholic acid

CN Ursodesoxycholic acid

CN Ursodiol

FS STEREOSEARCH

DR 50809-41-1, 80225-86-1

MF C24 H40 O4

CI COM

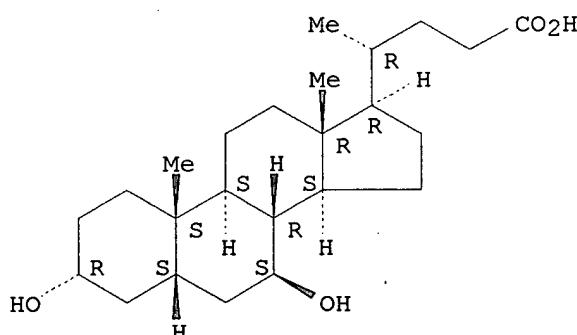
LC STN Files: ADISINSIGHT, AGRICOLA, AIDSLINE, ANABSTR, BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CABAB, CANCERLIT, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CBNB, CIN, CSCHEM, DETHERM\*, DDFU, DRUGU, EMBASE, HODOC\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, NAPRALERT, NIOSHTIC, PHAR, PROMT, RTECS\*, SPECINFO, TOXLINE, TOXLIT, USAN, USPATFULL, VETU

(\*File contains numerically searchable property data)

Other Sources: EINECS\*\*, WHO

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.



1818 REFERENCES IN FILE CA (1967 TO DATE)

67 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

1822 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
9 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 131:13243  
REFERENCE 2: 131:681  
REFERENCE 3: 130:350237  
REFERENCE 4: 130:332857  
REFERENCE 5: 130:322477  
REFERENCE 6: 130:308722  
REFERENCE 7: 130:306571  
REFERENCE 8: 130:306398  
REFERENCE 9: 130:301714  
REFERENCE 10: 130:278308

L105 ANSWER 8 OF 10 REGISTRY COPYRIGHT 1999 ACS

RN 81-23-2 REGISTRY

CN Cholan-24-oic acid, 3,7,12-trioxo-, (5.beta.)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5.beta.-Cholan-24-oic acid, 3,7,12-trioxo- (8CI)

OTHER NAMES:

CN 3,7,12-Triketo-5.beta.-cholanic acid

CN 3,7,12-Triketo-5.beta.-cholanoic acid

CN 3,7,12-Triketocholanic acid

CN 3,7,12-Trioxo-5.beta.-cholan-24-oic acid

CN 3,7,12-Trioxo-5.beta.-cholanic acid

CN 3,7,12-Trioxocholanic acid

CN Acolon

CN Bilioren

CN Bilstat

CN Cholagon

CN Cholic acid, dehydro-

CN Cholimed

CN Chologon

CN Decholin

CN Dehychol

CN Dehycon

CN Dehydrocholic acid

CN Dehystolin

CN Deidrocolico Vita

CN DHC

CN Didocol

CN Didrocolo

CN Dilabil

CN Drenobyl

CN Erebile

CN Felacrinos

CN Hydrochol

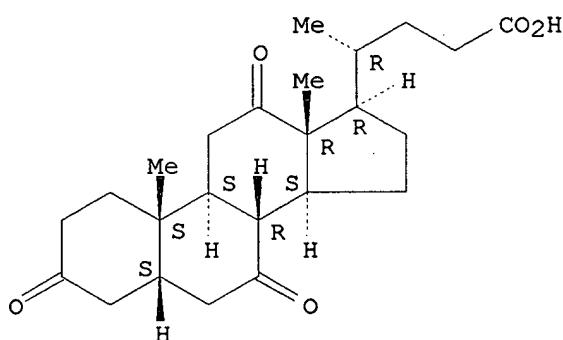
CN Hykolex

CN Ketocholanic acid

CN Novocolin

CN Oxycholin  
 CN Procholon  
 CN Sanocholen  
 CN Triketocholanic acid  
 FS STEREOSEARCH  
 MF C24 H34 O5  
 CI COM  
 LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CABA,  
 CANCERLIT, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMLIST, CSCHEM, DDFU,  
 DRUGU, EMBASE, HODOC\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*,  
 NAPRALERT, NIOSHTIC, PROMT, RTECS\*, SPECINFO, TOXLINE, TOXLIT, USAN,  
 USPATFULL, VETU  
 (\*File contains numerically searchable property data)  
 Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\*, WHO  
 (\*\*Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (+).



389 REFERENCES IN FILE CA (1967 TO DATE)  
 8 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 389 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
 5 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

- REFERENCE 1: 130:326572  
 REFERENCE 2: 130:322477  
 REFERENCE 3: 129:285538  
 REFERENCE 4: 129:270624  
 REFERENCE 5: 129:189522  
 REFERENCE 6: 129:8597  
 REFERENCE 7: 128:312903  
 REFERENCE 8: 128:288640  
 REFERENCE 9: 128:229877  
 REFERENCE 10: 128:229384

CN Cholest-5-en-3-ol (3. $\beta$ .)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Cholesterol (8CI)

OTHER NAMES:

CN (-)-Cholesterol

CN . $\Delta$ .5-Cholesten-3. $\beta$ .-ol

CN 3. $\beta$ .-Hydroxycholest-5-ene

CN 5:6-Cholesten-3. $\beta$ .-ol

CN Cholest-5-en-3. $\beta$ .-ol

CN Cholesterin

CN Cholesteryl alcohol

CN Dythol

CN Lidinit

CN Lidinite

CN Provitamin D

FS STEREOSEARCH

DR 209124-38-9, 218965-24-3

MF C27 H46 O

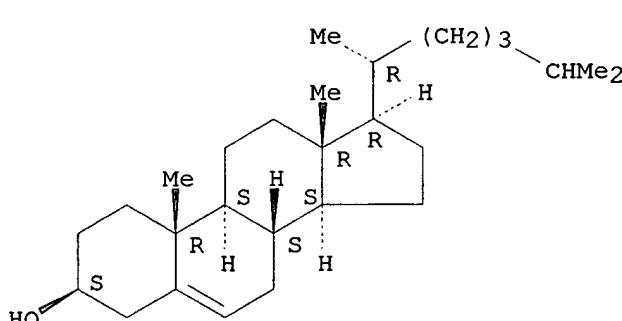
CI COM

LC STN Files: AGRICOLA, AIDSLINE, ANABSTR, BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CBNB, CIN, CSCHEM, CSNB, DETHERM\*, DDFU, DIPPR\*, DRUGU, EMBASE, GMELIN\*, HODOC\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, NAPRALERT, NIOSHTIC, PDLCOM\*, PIRA, PROMT, RTECS\*, SPECINFO, TOXLINE, TOXLIT, TULSA, ULIDAT, USAN, USPATFULL, VETU, VTB  
 (\*File contains numerically searchable property data)

Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.



65406 REFERENCES IN FILE CA (1967 TO DATE)

7293 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

65449 REFERENCES IN FILE CAPLUS (1967 TO DATE)

15 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 131:24646

REFERENCE 2: 131:23551

REFERENCE 3: 131:23549

REFERENCE 4: 131:23545

REFERENCE 5: 131:23536

REFERENCE 6: 131:23525

REFERENCE 7: 131:23407

REFERENCE 8: 131:23384

REFERENCE 9: 131:23340

REFERENCE 10: 131:23000

L105 ANSWER 10 OF 10 REGISTRY COPYRIGHT 1999 ACS

RN 57-10-3 REGISTRY

CN Hexadecanoic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Palmitic acid (7CI, 8CI)

OTHER NAMES:

CN 1-Pentadecanecarboxylic acid

CN Cetyllic acid

CN Emersol 143

CN Hydrofol Acid 1690

CN Hystrene 9016

CN Kortacid 1698

CN Loxiol EP 278

CN Lunac P 95

CN Lunac P 95KC

CN n-Hexadecanoic acid

CN n-Hexadecoic acid

CN NAA 160

CN Neo-Fat 16

CN PA 900

CN Palmitinic acid

CN Pentadecanecarboxylic acid

CN Prifac 2960

FS 3D CONCORD

DR 60605-23-4, 66321-94-6, 116860-99-2, 212625-86-0

MF C16 H32 O2

CI COM

LC STN Files: AGRICOLA, AIDSLINE, ANABSTR, APILIT, APILIT2, APIPAT,  
 APIPAT2, BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CABA, CANCERLIT, CAOLD,  
 CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CBNB, CIN, CSCHEM,  
 CSNB, DETHERM\*, DDFU, DIPPR\*, DRUGU, EMBASE, GMELIN\*, HODOC\*, HSDB\*,  
 IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, NAPRALERT,  
 NIOSHTIC, PDLCOM\*, PIRA, PROMT, RTECS\*, SPECINFO, TOXLINE, TOXLIT,  
 TRCTHERMO\*, TULSA, ULIDAT, USPATFULL, VETU, VTB

(\*File contains numerically searchable property data)

Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

HO2C-(CH2)14-Me

24288 REFERENCES IN FILE CA (1967 TO DATE)

927 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

24307 REFERENCES IN FILE CAPLUS (1967 TO DATE)

1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 131:23531

REFERENCE 2: 131:23496

REFERENCE 3: 131:23322

REFERENCE 4: 131:23217

REFERENCE 5: 131:22112

REFERENCE 6: 131:21236

REFERENCE 7: 131:20973

REFERENCE 8: 131:18443

REFERENCE 9: 131:18402

REFERENCE 10: 131:18392

=> fil hcaplus

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FILE COVERS 1967 - 6 Jul 1999 VOL 131 ISS 2  
FILE LAST UPDATED: 5 Jul 1999 (19990705/ED)

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=> d all 18

L8 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 1999 ACS  
AN 1998:765663 HCAPLUS  
DN 130:114999  
TI Preparation of slightly hydrophobic heparin derivatives which can be used for solvent casting in polymeric formulation  
AU Lee, Yong-Kyu; Moon, Hyun Tae; Byun, Youngro  
CS Department of Materials Science and Engineering, Kwangju Institute of Science and Technology, Kwangju, 506-712, S. Korea  
SO Thromb. Res. (1998), 92(4), 149-156  
CODEN: THBRAA; ISSN: 0049-3848  
PB Elsevier Science Inc.  
DT Journal  
LA English  
CC 63-8 (Pharmaceuticals)

AB **Heparin** is clin. administered mainly by i.v. injection because of its highly hydrophilic property. A slightly hydrophobic **heparin** deriv. which can be dissolved in org. solvent can be widely used in polymeric devices for clin. applications. In this study, hydrophobic **heparin** derivs. were prep'd. by coupling **heparin** with deoxycholic acid, cholesterol, lauric acid, and palmitic acid, resp. The hydrophobicity of these **heparin** derivs. depended on the feed mole ratio of **heparin** to hydrophobic agents, and they showed good solv. in the co-solvent of acetone and water, as well as in water alone. Also, these **heparin** derivs. showed high anticoagulant activity. This approach for prep'd. hydrophobic **heparin** is expected to advance the drug delivery system by further extending the applications of **heparin** to medical devices such as cardiopulmonary bypass circuits, heart lung oxygenators, and kidney dialyzers.

ST **heparin** hydrophobic deriv medical polymer

IT Antithrombogenic medical goods  
Hydrophobicity  
(prep'n. of slightly hydrophobic **heparin** derivs. which can be used for solvent casting in polymeric formulation)

IT 57-10-3DP, Palmitic acid, reaction products with **heparin**  
57-88-5DP, Cholesterol, reaction products with **heparin**  
83-44-3DP, Deoxycholic acid, reaction products with **heparin**  
143-07-7DP, Lauric acid, reaction products with **heparin**  
9005-49-6DP, **Heparin**, reaction products with cholates and alkanoic acids  
RL: BAC (Biological activity or effector, except adverse); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prep'n. of slightly hydrophobic **heparin** derivs. which can be used for solvent casting in polymeric formulation)

=> d bib abs hitrn tot 1101

L101 ANSWER 1 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
AN 1998:414634 HCAPLUS  
DN 129:72224  
TI Oral drug delivery compositions and methods  
IN Milstein, Sam J.; Barantsevitch, Evgueni N.; Sarubbi, Donald J.; Leone-Bay, Andrea; Paton, Duncan R.

PA Emisphere Technologies, Inc., USA  
SO U.S., 40 pp. Cont.-in-part of U.S. 5,451,410.  
CODEN: USXXAM

DT Patent  
LA English

FAN.CNT 17

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5766633	A	19980616	US 95-537888	19951023
	US 5451410	A	19950919	US 93-51019	19930422
	US 5792451	A	19980811	US 94-205511	19940302
PRAI	US 93-51019		19930422		
	US 94-205511		19940302		
	WO 94-US4560		19940422		

OS MARPAT 129:72224

AB The present invention relates to an oral drug delivery system, and in particular to modified amino acids and modified amino acid derivs. for use as a delivery system of sensitive agents such as bioactive peptides. The

modified amino acids and derivs. can form non-covalent mixts. with active biol. agents and in an alternate embodiment can releasably carry active agents. Modified amino acids can also form drug contg. microspheres. These mixts. are suitable for oral administration of biol. active agents to animals. Methods for the prepn. of such amino acids are also disclosed. In a test tube 568 mg acetyl phenylalanine aldehyde, 132 mg carbomethoxyphenylalanylleucine, and 100 mg N-acetyl-Phe-Leu-Leu-Arg aldehyde were added to 2.9 mL of 15 % ethanol. The soln. was stirred and NaOH was added to raise the pH to 7.2 and water was added to bring the total vol. to 4 mL. Calcitonin 6 .mu.g was added to the soln. to obtain an oral soln.

IT 9005-49-6, Heparin, biological studies  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (amino acid derivs. for oral delivery of sensitive biol. active agents)

L101 ANSWER 2 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1998:195043 HCPLUS  
 DN 128:248635  
 TI Small caliber vascular grafts with significant patency enhancement via a surface coating which contains **covalently** bonded bioactive agents  
 IN Patnaik, Birendra K.; Lin, Horng-Ban; Lentz, David J.; Zdrhala, Richard J.  
 PA Meadox Medicals, Inc., USA  
 SO PCT Int. Appl., 33 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9810806	A1	19980319	WO 97-US16163	19970911
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	AU 9743436	A1	19980402	AU 97-43436	19970911
PRAI	US 96-713800		19960913		
	WO 97-US16163		19970911		
AB	Disclosed are implantable medical devices with enhanced patency. Expanded polytetrafluoroethylene small caliber vascular grafts coated with polymer bound bio-active agents that exhibit enhanced patency are disclosed. In a preferred embodiment, prior to deposition of the bioactive coating, the <b>hydrophobic</b> surface is treated with a hydrogen-rich plasma to provide better adhesion of the coating. The plasma-primed ePTFE grafts are then contacted with coating materials, which comprise bioactive agents, such as heparin, <b>covalently</b> bound to a polymer-spacer complex in the presence of a dehydrating agent.				
IT	9005-49-6, Heparin, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (small caliber vascular grafts coated with polymer-bound bioactive agents)				

L101 ANSWER 3 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1997:503011 HCPLUS

DN 127:113410  
 TI Crosslinked polymer compositions and methods for their use  
 IN Rhee, Woonza M.; Delustro, Frank A.; Berg, Richard A.  
 PA Collagen Corporation, USA  
 SO PCT Int. Appl., 75 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9722371	A1	19970626	WO 96-US19975	19961218
	W: AU, CA, JP				
	RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	AU 9713344	A1	19970714	AU 97-13344	19961218
	EP 876165	A1	19981111	EP 96-944824	19961218
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
PRAI	US 95-573799		19951218		
	WO 96-US19975		19961218		
AB	Crosslinked polymer compns. comprise a first synthetic polymer contg. multiple nucleophilic groups <b>covalently</b> bound to a second synthetic polymer contg. multiple electrophilic groups. The first synthetic polymer is preferably a synthetic polypeptide or a polyethylene glycol that has been modified to contain multiple nucleophilic groups, such as primary amino (-NH <sub>2</sub> ) or thiol (-SH) groups. The second synthetic polymer may be a hydrophilic or <b>hydrophobic</b> synthetic polymer which contains, or has been derivatized to contain, two or more electrophilic groups, such as succinimidyl groups. The compns. may further comprise other components, such as naturally occurring polysaccharides or proteins (such as glycosaminoglycans or collagen) and/or biol. active agents. Also disclosed are methods for using the crosslinked polymer compns. to effect adhesion between a first surface and a second surface; to effect tissue augmentation; to prevent the formation of surgical adhesions; and to coat a surface of a synthetic implant, such as those for cell delivery.				
IT	<b>9005-49-6</b> , Heparin, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (in crosslinked polymer compns. for prosthetic implants)				

L101 ANSWER 4 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1997:379803 HCPLUS  
 DN 127:86043  
 TI Covalent bonding of heparin to a vinyl copolymer for biomedical applications  
 AU Marconi, W.; Benvenuti, F.; Piozzi, A.  
 CS Dep. Chem., Univ. Rome "La Sapienza", Rome, 00185, Italy  
 SO Biomaterials (1997), 18(12), 885-890  
 CODEN: BIMADU; ISSN: 0142-9612  
 PB Elsevier  
 DT Journal  
 LA English  
 AB In order to prep. polymer surfaces of vinyl type, provided with long-term hemocompatibility, a com. ethylene-vinyl alc. copolymer (EVAL) was **covalently** heparinized, employing two different bifunctional reagents (adipoyl chloride and hexamethylene diisocyanate). The amt. and activity of the heparin bonded to the polymer films were evaluated as a function of the concn. of the heparin solns. employed. Also, the influence exerted by the presence of various hydrophilic 'spacing arms' of

different mol. wts., either neutral or provided with elec. charge, was investigated. By in vitro measurements of activated partial thromboplastin time it was seen that all the heparinized samples possessed a high degree of hemocompatibility. For the sake of comparison, heparin was also bonded ionically to EVAL functionalized by introduction of quaternary ammonium groups bonded covalently (by adipoyl chloride) to the hydroxyl groups of the polymer. It was seen that the covalent immobilization system provides the polymer surfaces with a superior hemocompatibility.

IT 9005-49-6DP, Heparin, reaction products EVA 24937-78-8DP  
, Eva, reaction products with spacers and heparin  
RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);  
BIOL (Biological study); PREP (Preparation); USES (Uses)  
(covalent bonding of heparin to a vinyl copolymer for  
biomedical applications)

L101 ANSWER 5 OF 15 HCPLUS COPYRIGHT 1999 ACS

AN 1997:224081 HCPLUS

DN 126:216675

TI Drug delivery systems for macromolecular drugs

IN Dunn, James M.

PA Dunn, James M., USA

SO PCT Int. Appl., 24 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9704747	A1	19970213	WO 96-US12203	19960725
	W: CA, JP, MX				
	RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				

PRAI US 95-508247 19950727

AB This invention described the oral, parenteral or inhalation delivery of large macromols. Biol. active drugs are entrapped into biodegradable hydrogel polymers in either org. and water phase systems. By using cyclodextrins, sensitive mols. can be protected during the granulations of nanoparticles prodn. phase. Aq. insulin (100 mL; 10,000U) was mixed with 100 g .beta.-cyclodextrin and blended to give a clear soln. To this soln. was added 500 mg liposomal Phospholipon 90H and the soln. blended, then 1000 mg a 30% dispersion of Me methacrylate was added. The mixt. was dried at 60.degree. for 4-6 h until dry. The effectiveness of the insulin nanoparticles in decreasing blood glucose levels was demonstrated in rats.

IT 9005-49-6, Heparin, biological studies 9041-08-1,

Heparin sodium

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(polymeric drug delivery systems for macromol. drugs)

IT 57-88-5, Cholesterol, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(polymeric drug delivery systems for macromol. drugs)

L101 ANSWER 6 OF 15 HCPLUS COPYRIGHT 1999 ACS

AN 1996:544101 HCPLUS

DN 125:177462

TI Surface-modified nanoparticles and method of making and using them

IN Levy, Robert J.; Labhasetwar, Vinod; Song, Cunxian S.

PA USA

SO PCT Int. Appl., 170 pp.

CODEN: PIXXD2

DT Patent  
LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9620698	A2	19960711	WO 96-US476	19960104
	WO 9620698	A3	19980122		
	W: AL, AM, AT, AU, CA, CH, CN, CZ, DE, DK, GB, HU, IS, JP, KE, LU, VN, MN, NO, US				
	RW: KE, LS, SD, AT, BE, CH, DE, ES, FR, GB, IT, LU, NL, PT, SE, NL, MR, NE, SN				
	CA 2207961	AA	19960711	CA 96-2207961	19960104
	AU 9647556	A1	19960724	AU 96-47556	19960104
	EP 805678	A1	19971112	EP 96-903476	19960104
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE				
	JP 10511957	T2	19981117	JP 96-521279	19960104
PRAI	US 95-369541		19950105		
	US 95-389893		19950216		
	WO 96-US476		19960104		
AB	Biodegradable controlled-release nanoparticles as sustained release bioactive agent delivery vehicles include surface modifying agents to target binding of the nanoparticles to tissues or cells of living systems, to enhance nanoparticle sustained release properties, and to protect nanoparticle-incorporated bioactive agents. Unique methods of making small (10 nm to 15 nm, and preferably 20 nm to 35 nm) nanoparticles having a narrow size distribution which can be surface-modified after the nanoparticles are formed is described. Techniques for modifying the surface include a lyophilization technique to produce a phys. adsorbed coating and epoxy-derivatization to functionalize the surface of the nanoparticles to covalently bind mols. of interest. The nanoparticles may also comprise hydroxy-terminated or epoxide-terminated and/or activated multiblock copolymers, having hydrophobic segments which may be polycaprolactone and hydrophilic segments. The nanoparticles are useful for local intravascular administration of smooth muscle inhibitors and antithrombogenic agents as part of interventional cardiac or vascular catheterization such as a balloon angioplasty procedure; direct application to tissues and/or cells for gene therapy, such as the delivery of osteotropic genes or gene segments into bone progenitor cells; or oral administration in an enteric capsule for delivery of protein/peptide based vaccines.				
IT	57-10-3, Hexadecanoic acid, biological studies 57-88-5, Cholesterol, biological studies 9005-49-6, Heparin, biological studies				
	RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(surface-modified polymer controlled-release nanoparticles for sustained drug delivery)				
IT	9005-49-6DP, Heparin, reaction products with epoxide end-capped polymer				
	RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)				
	(surface-modified polymer controlled-release nanoparticles for sustained drug delivery)				

TI Study of hemodialysis materials: physicochemical and biological characterization of EVALVA, EVAPA, and heparinized EVAPA  
 AU Barbucci, R.; Albanese, A.; Tempesti, F.; Baszkin, A.; Eloy, R.; Weill, N.; Martuscelli, E.; Cimmino, S.  
 CS CRISMA, Universita di Siena, Siena, 53100, Italy  
 SO J. Mater. Sci.: Mater. Med. (1994), 5(12), 844-9  
 CODEN: JSMMEL; ISSN: 0957-4530  
 DT Journal  
 LA English  
 AB Partially hydrolyzed ethylene/vinyl acetate copolymers were modified by the covalent binding of a heparin-complexing polymer and further heparinized in order to improve their blood compatibility. These heparinizable polymeric materials (EVAPA) were obtained by a 2-step reaction between an ethylene/vinyl alc./vinyl acetate (EVALVA) terpolymer, and the heparin complexing polymer N2LL. The physicochem. characterization of EVALVA, EVAPA and heparinized-EVAPA was carried out through thermal anal., SEM, contact angle, potentiometric measurements, water uptake and FT-IR spectroscopic measurements. The biocompatibility of the above-mentioned samples was evaluated using in vitro methods, through the detn. of heparin release in phosphate buffer soln. and in human plasma, and with the investigation of hemostasis activation.  
 IT 9005-49-6D, Heparin, reaction products with polyamide-polyamines and ethylene-vinyl alc.-vinyl acetate copolymer 24937-78-8D, Ethylene-vinyl acetate copolymer, hydrolyzed, reaction products with polyamide-polyamines  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (physicochem. and biol. characterization of heparinized hemodialysis polymers)

L101 ANSWER 8 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
 AN 1994:226984 HCAPLUS  
 DN 120:226984  
 TI Compositions of oral nondissolvable matrixes for transmucosal administration of medicaments  
 IN Stanley, Theodore H.; Hague, Brian  
 PA University of Utah Research Foundation, USA  
 SO U.S., 20 pp. Cont.-in-part of U.S. 4,863,737.  
 CODEN: USXXAM

DT Patent  
 LA English  
 FAN.CNT 9

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5288498	A	19940222	US 89-403752	19890905
	US 4671953	A	19870609	US 85-729301	19850501
	JP 05501539	T2	19930325	JP 89-504878	19890816
	JP 2801050	B2	19980921		
	AU 641127	B2	19930916	AU 89-40704	19890816
	EP 487520	B1	19950412	EP 89-909497	19890816
	R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE				
	AT 120953	E	19950415	AT 89-909497	19890816
	CA 1338978	A1	19970311	CA 89-609378	19890824
	AU 9050352	A1	19910408	AU 90-50352	19890905
	AU 645966	B2	19940203		
	EP 493380	A1	19920708	EP 90-902584	19890905
	EP 493380	B1	19971029		
	R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE				
	US 5132114	A	19920721	US 89-402881	19890905
	JP 05501854	T2	19930408	JP 90-502779	19890905

CA 1339075	A1	19970729	CA 89-610329	19890905
AT 159658	E	19971115	AT 90-902584	19890905
WO 9103236	A1	19910321	WO 90-US4369	19900803
W: AU, CA, JP, NO				
RW: AT, BE, CH, DE, DK, ES, FR, GB, IT, LU, NL, SE				
AU 9063371	A1	19910408	AU 90-63371	19900803
AU 642664	B2	19931028		
EP 490944	A1	19920624	EP 90-913359	19900803
EP 490944	B1	19960529		
R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE				
JP 05500058	T2	19930114	JP 90-512483	19900803
JP 2749198	B2	19980513		
AT 138562	E	19960615	AT 90-913359	19900803
ES 2089027	T3	19961001	ES 90-913359	19900803
CA 2066403	C	19980414	CA 90-2066403	19900803
NO 9200565	A	19920213	NO 92-565	19920213
DK 9200193	A	19920214	DK 92-193	19920214
NO 9200858	A	19920304	NO 92-858	19920304
NO 9200855	A	19920410	NO 92-855	19920304
NO 9200854	A	19920427	NO 92-854	19920304
DK 9200300	A	19920505	DK 92-300	19920305
AU 9460697	A1	19940623	AU 94-60697	19940427
US 5855908	A	19990105	US 94-339655	19941115

PRAI US 85-729301 19850501  
 US 87-60045 19870608  
 EP 89-909497 19890816  
 WO 89-US3518 19890816  
 US 89-403752 19890905  
 WO 89-US3801 19890905  
 WO 90-US4369 19900803  
 US 93-152414 19931112

AB Compns. and methods of manuf. for producing a medicament compn. capable of absorption through the mucosal tissues of the mouth, pharynx, and esophagus are disclosed. The present invention relates to such compns. and methods which are useful in administering lipophilic and nonlipophilic drugs in a dose-to-effect manner such that sufficient drug is administered to produce precisely a desired effect. The invention also relates to manufg. techniques that enable therapeutic agents to be incorporated into nondissolvable drug containment matrixes which are capable of releasing the drug within a patient's mouth. An appliance or holder is preferably attached to the drug containment matrix. Employing the present invention the drug may be introduced into the patient's bloodstream almost as fast as through injection, and much faster than using the oral administration route, while avoiding the neg. aspects of both of these methods. The nondissolvable drug containment matrix may include permeation enhancers to increase the drug adsorption by the mucosal tissues of the mouth. The matrix compn. may also include pH buffering agents to modify the saliva pH thereby increasing the absorption of the drug through the mucosal tissues. Figures show views of some dosage forms.

IT 81-23-2, Dehydrocholate 128-13-2, Ursodeoxycholate  
 474-25-9, Chenodeoxycholate 2955-27-3, Ursocholate  
 RL: BIOL (Biological study)

(as permeation enhancer for transmucosal pharmaceuticals)

IT 9005-49-6, Heparin, biological studies  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (transmucosal pharmaceuticals contg.)

DN 120:226981  
 TI Compositions of oral dissolvable medicaments  
 IN Stanley, Theodore H.; Hague, Brian  
 PA University of Utah, USA  
 SO U.S., 22 pp. Cont.-in-part of U.S. 4,863,737.  
 CODEN: USXXAM  
 DT Patent  
 LA English  
 FAN.CNT 9

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5288497	A	19940222	US 89-403751	19890905
	US 4671953	A	19870609	US 85-729301	19850501
	JP 05501539	T2	19930325	JP 89-504878	19890816
	JP 2801050	B2	19980921		
	AU 641127	B2	19930916	AU 89-40704	19890816
	EP 487520	B1	19950412	EP 89-909497	19890816
	R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE				
	AT 120953	E	19950415	AT 89-909497	19890816
	CA 1338978	A1	19970311	CA 89-609378	19890824
	AU 9050352	A1	19910408	AU 90-50352	19890905
AU 645966	B2	19940203			
EP 493380	A1	19920708	EP 90-902584	19890905	
EP 493380	B1	19971029			
R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE					
US 5132114	A	19920721	US 89-402881	19890905	
JP 05501854	T2	19930408	JP 90-502779	19890905	
CA 1339075	A1	19970729	CA 89-610329	19890905	
AT 159658	E	19971115	AT 90-902584	19890905	
WO 9103237	A1	19910321	WO 90-US4384	19900803	
W: AU, CA, JP, NO					
RW: AT, BE, CH, DE, DK, ES, FR, GB, IT, LU, NL, SE					
AU 9062877	A1	19910408	AU 90-62877	19900803	
AU 645265	B2	19940113			
EP 490916	A1	19920624	EP 90-912733	19900803	
EP 490916	B1	19951018			
R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE					
JP 05503917	T2	19930624	JP 90-512229	19900803	
EP 630647	A1	19941228	EP 94-111352	19900803	
EP 630647	B1	19990303			
R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE					
AT 129148	E	19951115	AT 90-912733	19900803	
ES 2077686	T3	19951201	ES 90-912733	19900803	
CA 2066423	C	19980414	CA 90-2066423	19900803	
AT 177007	E	19990315	AT 94-111352	19900803	
NO 9200565	A	19920213	NO 92-565	19920213	
DK 9200193	A	19920214	DK 92-193	19920214	
NO 9200857	A	19920406	NO 92-857	19920304	
NO 9200855	A	19920410	NO 92-855	19920304	
NO 9200854	A	19920427	NO 92-854	19920304	
DK 9200300	A	19920505	DK 92-300	19920305	
AU 9455218	A1	19940428	AU 94-55218	19940218	
AU 668004	B2	19960418			
AU 9460697	A1	19940623	AU 94-60697	19940427	
US 5824334	A	19981020	US 96-636828	19960419	
US 5783207	A	19980721	US 97-795359	19970204	
US 5785989	A	19980728	US 97-822560	19970319	
PRAI US 85-729301		19850501			
US 87-60045		19870608			

EP 89-909497	19890816
WO 89-US3518	19890816
US 89-403751	19890905
WO 89-US3801	19890905
EP 90-912733	19900803
WO 90-US4384	19900803
US 93-152396	19931112
US 94-333233	19941102
US 95-439127	19950511

AB Compns. and methods of manuf. for producing a medicament compn. capable of absorption through the mucosal tissues of the mouth, pharynx, and esophagus are disclosed. The present invention relates to such compns. and methods which are useful in administering lipophilic and nonlipophilic drugs in a dose-to-effect manner that sufficient drug is administered to produce precisely a desired effect. The invention also relates to a manufg. technique that enables a therapeutic agent or drug to be incorporated into a flavored dissolvable matrix. An appliance or holder is preferably attached to the dissolvable matrix. Employing the present invention the drug may be introduced into the patient's bloodstream almost as fast as through injection, and much faster than using the oral administration route, while avoiding the neg. aspects of both of these methods. The present invention achieves these advantages by incorporating the drug into a carbohydrate, fat, protein, wax, or other dissolvable matrix compn. The dissolvable matrix may include permeation enhancers to increase the drug absorption by the mucosal tissues of the mouth. The matrix compn. may also include pH buffering agents to modify the salival pH thereby increasing the absorption of the drug through the mucosal tissue. Methohexital sodium was incorporated into a dissolvable matrix including citric acid; ribotide; Compritol 888; aspartame; vanilla, wild cherry, and peppermint microcapsules; compressible sugar; and maltodextrin.

IT 81-23-2, Dehydrocholate 128-13-2, Ursodeoxycholate  
 474-25-9, Chenodeoxycholate 2955-27-3, Ursocholate  
 RL: BIOL (Biological study)  
 (as permeation enhancer for transmucosal pharmaceuticals)

IT 9005-49-6, Heparin, biological studies  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (transmucosal pharmaceuticals contg.)

L101 ANSWER 10 OF 15 HCAPLUS COPYRIGHT 1999 ACS

AN 1994:200438 HCAPLUS

DN 120:200438

TI Controlled-release transdermal pharmaceuticals  
 containing cyrogels

IN Wood, Louis L.; Calton, Gary J.

PA SRCHEM Inc., USA

SO U.S., 15 pp.

CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5260066	A	19931109	US 92-821627	19920116
	US 5288503	A	19940222	US 92-899369	19920616

PRAI US 92-821627 19920116

AB A controlled-release transdermal pharmaceutical contg.  
 therapeutic agents in a poly(vinyl alc.) (I) cyrogel is disclosed. A slurry of 11.0 mg ciprofloxacin.HCl (II) and 200 mg 10% I was warmed to

50-60.degree. to obtain a clear homogeneous soln. The soln. was then placed in a mold and subjected to 6 freeze-thaw cycles to give a white opaque elastomeric cryogel having 15mm diam. and 0.5mm thickness. The release of II from the gel in 0.9% NaCl was 74% in th 1st 4 hs and it was const. in the subsequent 5-24 hs.

IT 81-23-2, Dehydrocholic acid 9005-49-6, Heparin,  
biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(controlled-release transdermal pharmaceuticals  
contg. cryogels and)

L101 ANSWER 11 OF 15 HCPLUS COPYRIGHT 1999 ACS

AN 1989:484146 HCPLUS

DN 111:84146

TI Polymeric articles having antithrombogenic activity

IN Hu, Can B.; Solomon, Donald D.

PA Becton, Dickinson and Co., USA

SO U.S., 10 pp.

CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 4720512	A	19880119	US 86-843323	19860324
	US 4786556	A	19881122	US 87-77468	19870724

PRAI US 86-843323 19860324

AB Plastic articles are prep'd. which have enhanced antithrombogenic activity due to the presence of hydrophobic materials attached to amine compds. bonded on the surface of the plastic which repel bonded antithrombogenic agents outwardly from the plastic surface, making them more readily available to the blood for subsequent thrombus inhibition. The amine-rich surface is plasma-treated with a gaseous material in an ionization chamber whereby hydrophobic groups generated from the gaseous material are introduced to give a hydrophobic amine-rich surface. Trimethylolpropane (44.73 g) and 337.59 g Terathane 650 (low-mol.-wt. polyether-polyol) (1 equiv each) were mixed at 45.degree., 524.00 g (4 equiv) hydrogenated diphenylmethane diisocyanate and 14 g (0.015%) dibutyltin dilaurate were added and after 5 min of mixing, the reactants were transferred to a 90.degree. oven for 60 min to give a prepolymer. This prepolymer (15 g) was added to 30 g toluene to make a 33% wt./wt. soln. This prepolymer soln. was added dropwise (2 h) to a sep. diamine soln. contg. 4.1 g 1,6-hexanediamine, 20 g iso-PrOH and 10 g toluene and stirred an addnl. 2 h to give an amine-rich polyurethane-urea (APU). This was dissolved in MeOH to a 20% wt./wt. soln. for deposition on the polymeric support structure. Aldehyde-activated heparin for bonding to the amine compd. was prep'd. by the periodate oxidn. of heparin. Polyurethane tubing was coated with the APU soln. and dried in a N atm. at ambient temp. for 60 min. The coated tubing was then treated with hexafluoropropylene plasma (0.3 torr and 50 W) for 15 min. Samples were then treated with 2% aq. aldehyde-activated heparin (pH 6.6, 10N NaOH) and 0.025 g Na cyanoborohydride at 50.degree. for 2 h, washed with 3M saline for 1 h, and treated for a 2nd 2 h period at the same conditions to give a material that had 87.8 .mu.g heparin bonded (radiolabel assays) per cm<sup>2</sup> of surface area compared with 143.4 .mu.g/cm<sup>2</sup> for control samples which were identically treated except without plasma treatment. After a dynamic leach study in 3M saline for 24 h, 79.4 .mu.g heparin/cm<sup>2</sup> remained on the plasma-treated sample compared with 103.7 .mu.g/cm<sup>2</sup> for the control demonstrating the permanency of the covalently bonded heparin.

The increased antithrombogenicity of a plasma-treated sample was demonstrated in an in vivo study by inserting it in 1 external jugular vein of a same animal and measuring platelet adhesion (platelet uptake slope) and thrombus wt. which were 0.0188 and 4.53 mg, resp., for the plasma-treated sample compared with 0.0413 and 14.13 mg, resp., for the control sample; the lower slope and lower thrombus wt. indicated increased antithrombogenic activity.

IT 9005-49-6D, Heparin, reaction products with hydrophobic amine-rich polymeric substrate  
 RL: BIOL (Biological study)  
 (as antithrombogenic material)

L101 ANSWER 12 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1988:156424 HCPLUS  
 DN 108:156424  
 TI Covalent immobilization of chitosan derivatives onto polymeric film surfaces with the use of a photosensitive hetero-bifunctional crosslinking reagent  
 AU Aiba, Seiichi; Minoura, Norihiko; Taguchi, Kazuhiro; Fujiwara, Yukihiko  
 CS Funct. Polym. Div., Ind. Prod. Res. Inst., Yatabe, 305, Japan  
 SO Biomaterials (1987), 8(6), 481-8  
 CODEN: BIMADU; ISSN: 0142-9612  
 DT Journal  
 LA English  
 AB Partially N-acetylated chitosan was covalently immobilized onto polymeric film surfaces using the photosensitive heterobifunctional crosslinking reagent, Me 4-azidobenzimidate, which was previously attached to the chitosan by the reaction between an imidoester group of the reagent and a free amino group of the chitosan. The grafting was accomplished by irradiating with UV light, the modified chitosan being coated on the film surfaces to photolyze arylazide groups, thus crosslinking the chitosan and the underlying substrate polymer together. For UV absorption and IR spectroscopy, the irradn. time of 3 min was sufficient for the photolysis of the azide group. The thickness of the immobilized chitosan layer was estd. to be of the order of 30-150 nm using attenuated total reflectance Fourier transform IR spectroscopy (ATR-FTIR). The chitosan mols. immobilized on the surfaces could be chem. modified by several reagents and also treated with a heparin soln. to form a polyelectrolyte complex on the surface. The ionically bound heparin was partially released into a phosphate buffer soln.  
 IT 9005-49-6DP, Heparin, reaction products with chitosan derivs. bound on polymer surfaces 24937-78-8DP, Ethylene vinyl acetate copolymer, reaction products with crosslinked chitosan  
 RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and properties of)

L101 ANSWER 13 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1986:10617 HCPLUS  
 DN 104:10617  
 TI Liposomes containing pharmaceuticals  
 IN Honda, Haruo; Maruyama, Tetsupei; Saito, Noriko  
 PA Terumo Corp., Japan  
 SO Jpn. Kokai Tokkyo Koho, 6 pp.  
 CODEN: JKXXAF  
 DT Patent  
 LA Japanese  
 FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI JP 60155109 A2 19850815 JP 84-9910 19840123  
 JP 63054684 B4 19881028

AB Liposome-contg. drugs are prepd. for stabilization and sustained release of drugs. The liposome membranes are made of lipids and C10-20 fatty acids. Thus, lecithin 32, cholesterol [57-88-5] 15.4, and oleic acid [112-80-1] 5.0 mg were dissolved in 3 mL CHCl<sub>3</sub>, mixed with 3 mL iso-Pr ether and 1 mL H<sub>2</sub>O contg. 10,000 units urokinase [9039-53-6], treated with ultrasound 5 min at 4.degree., and dried by a rotary evaporator at 40.degree. to give a gel. The gel was mixed with 5 mL saline, and the solvent removed. The suspension was centrifuged at 100,000 times. g for 30 min. The sediment was washed with saline to give urokinase-contg. liposomes.

IT 544-63-8, biological studies

RL: BIOL (Biological study)  
 (liposomes contg. cholesterol and, for drug formulation)

IT 57-88-5, biological studies

RL: BIOL (Biological study)  
 (liposomes contg. fatty acids and, for drug formulation)

IT 9005-49-6, biological studies

RL: BIOL (Biological study)  
 (liposomes contg., for pharmaceuticals)

L101 ANSWER 14 OF 15 HCAPLUS COPYRIGHT 1999 ACS

AN 1985:100827 HCAPLUS

DN 102:100827

TI Liposome formulations containing pharmaceuticals

PA Terumo Corp., Japan

SO Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 59222410	A2	19841214	JP 83-95674	19830601

AB Sustained-release, stable pharmaceuticals consist of liposomes contg. drugs. The liposomes are made of lecithins or natural phospholipids contg. sterols such as cholesterol [57-88-5]. Thus, phospholipids, cholesterol, and dicetyl phosphate [2197-63-9] (7:7:1 mol ratio) were mixed and dissolved in CHCl<sub>3</sub>, and the soln. was dried in a rotating flask to form a film on the wall of the flask. The film was dissolved in a CHCl<sub>3</sub>-iso-Pr ether (1:1) mixt., mixed with drugs, treated with ultrasonic waves, and dried in a rotary evaporator to obtain liposomes.

IT 57-88-5, biological studies

RL: BIOL (Biological study)  
 (liposomes contg. phospholipids and, for sustained-release pharmaceuticals)

IT 9005-49-6, biological studies

RL: BIOL (Biological study)  
 (sustained-release pharmaceuticals contg. liposomes and)

L101 ANSWER 15 OF 15 HCAPLUS COPYRIGHT 1999 ACS

AN 1982:461071 HCAPLUS

DN 97:61071

TI Nonthrombogenic material comprising substrate which has been reacted with heparin

IN Joh, Yasushi

PA Nippon Zeon Co., Ltd. , Japan  
 SO U.S., 10 pp. Division of U.S. Ser. No. 60,054.  
 CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 4329383	A	19820511	US 80-170656	19800721
	US 4415490	A	19831115	US 79-60054	19790724

PRAI US 79-60054 19790724

AB Nonthrombogenic materials for blood contacting medical devices such as extracorporeal systems are prep'd. by covalently linking heparin with base aldehyde group-contg. polymers through only 1 acetal or hemiacetal bond at each bonding site. Polymers from aldehyde group-contg. monomers or natural polymers with glucose units, treated with HIO4 which converts them to aldehyde group-contg. polymers are used. Examples included cuprophane and cellophane films, or starch treated with HIO4 and then heparin. Other polymers such as vinyl acetate-ethylene copolymer or allylidene diacetate-vinyl acetate copolymer were hydrolyzed and the former treated with Na heparin-NaIO4 and the latter with heparin. Coagulation times with fresh human blood were much greater for the heparinized polymers compared to nonheparinized ones or glass plate control.

IT 9005-49-6DP, reaction products with formaldehyde-group-contg. polymer 24937-78-8DP, hydrolyzed, metaperiodate oxidized, reaction products with heparin

RL: PREP (Preparation)  
 (prep'n. of, as antithrombogenic material)

=> d his 1106-

(FILE 'REGISTRY' ENTERED AT 12:24:33 ON 06 JUL 1999)

FILE 'HCAPLUS' ENTERED AT 12:25:22 ON 06 JUL 1999

E HYDROPHOB/CW

L106 5713 S E4-E6  
 L107 78 S L106 AND L51  
 L108 4 S L107 AND L60  
 L109 3 S L108 NOT L8,L101  
 L110 1 S L107 AND (BILE OR STEROL OR ALKANOIC)  
 L111 0 S L110 NOT L8  
 L112 13 S L107 AND 63/SC  
 L113 11 S L112 NOT L8,L109,L101  
 L114 15 S L109,L110,L113

=> d bib abs hitrn tot

L114 ANSWER 1 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
 AN 1998:765663 HCAPLUS

DN 130:114999

TI Preparation of slightly hydrophobic heparin derivatives which can be used for solvent casting in polymeric formulation

AU Lee, Yong-Kyu; Moon, Hyun Tae; Byun, Youngro

CS Department of Materials Science and Engineering, Kwangju Institute of Science and Technology, Kwangju, 506-712, S. Korea

SO Thromb. Res. (1998), 92(4), 149-156

CODEN: THBRAA; ISSN: 0049-3848

PB Elsevier Science Inc.

DT Journal

LA English

AB **Heparin** is clin. administered mainly by i.v. injection because of its highly hydrophilic property. A slightly hydrophobic **heparin** deriv. which can be dissolved in org. solvent can be widely used in polymeric devices for clin. applications. In this study, hydrophobic **heparin** derivs. were prep'd. by coupling **heparin** with deoxycholic acid, cholesterol, lauric acid, and palmitic acid, resp. The hydrophobicity of these **heparin** derivs. depended on the feed mole ratio of **heparin** to hydrophobic agents, and they showed good solv. in the co-solvent of acetone and water, as well as in water alone. Also, these **heparin** derivs. showed high anticoagulant activity. This approach for prep'd. hydrophobic **heparin** is expected to advance the drug delivery system by further extending the applications of **heparin** to medical devices such as cardiopulmonary bypass circuits, heart lung oxygenators, and kidney dialyzers.

L114 ANSWER 2 OF 15 HCAPLUS COPYRIGHT 1999 ACS

AN 1998:708924 HCAPLUS

DN 129:335768

TI Controlled release formulations using intelligent polymers

IN Odidi, Isa; Odidi, Amina

PA Can.

SO PCT Int. Appl., 24 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9847491	A2	19981029	WO 98-CA274	19980403
	WO 9847491	A3	19990121		
	W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			
	CA 2216215	AA	19981005	CA 97-2216215	19971117
	AU 9868170	A1	19981113	AU 98-68170	19980403

PRAI US 97-36551 19970421

WO 98-CA274 19980403

AB An extended release dosage compn. of pharmaceutically active substances that have a water contact angle (.theta.) such that cos .theta. is between +0.9848 and -0.9848 presented as a matrix tablet contg. the said pharmaceutically active substances, with/without suitable pharmaceutical excipients in intimate mixt. with two groups of intelligent polymers having opposing wettability characteristics, one demonstrating a stronger tendency towards hydrophobicity and the other a stronger tendency towards hydrophilicity, the polymer combination being between the ratios of 1:50 and 50:1 amts. effective to control the release of said pharmaceutically active substances in a math. predictable manner, wherein the polymer demonstrating a stronger tendency towards hydrophobicity is not less than 5 % wt/wt and preferably between 5-70 % wt/wt of the final formulation

comprn. The intelligent polymers being Et cellulose (EC) as a more strongly hydrophobic and hydroxyethyl cellulose (HEC) and/or hydroxypropyl Me cellulose (HPMC) as more strongly hydrophilic (the ratio of HEC to HPMC being between 1:100 and 100:1). The matrix tablet is optionally coated with an enteric coat, 0-5 % - 15 % wt/wt to prevent the initial burst effect seen in such systems and to impart gastrointestinal tract (GIT) "stealth" characteristics esp. in the presence of food. A comprn. was prep'd. contg. HPMC 20, glipizide 1.83, Et cellulose 16.17, hydroxyethyl cellulose 4, lactose 30, microcryst. cellulose 23, SiO<sub>2</sub> 0.6, Na lauryl sulfate 4, and Mg stearate 0.4%.

L114 ANSWER 3 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1998:190439 HCPLUS  
 DN 128:221553  
 TI The use of bead celluloses as carrier for controlled delivery of drugs. Part 10. Lipophilic coprecipitates of bead cellulose and fatty oils  
 AU Wolf, Bertram  
 CS Dep. Pharmaceutical Technol., Inst. Pharmacy, Fac. Biological Sciences, Pharmacy, Psychology, Univ. Leipzig, Leipzig, D-04207, Germany  
 SO Pharmazie (1998), 53(3), 177-180  
 CODEN: PHARAT; ISSN: 0031-7144  
 PB Govi-Verlag Pharmazeutischer Verlag  
 DT Journal  
 LA English  
 AB Coppts. of bead cellulose (BC) and strong lipophilic liqs. (white mineral oil (WMO), oenotheric oil, olive oil, juniper berry oil and peanut oil) were prep'd. with different wt. ratio. BC was able to take up an WMO amt. of max. 25% of cellulose dry wt. The product was flowable and consisted of spherical and not cohesive particles, higher WMO content yielded agglomerated products. WMO was incorporated into the pores of BC and also pptd. on the surface of the beads. The uptake of water by the product was low due to predominant hydrophobic properties. BC/ oil coppts. represent a suitable lipophilic drug carrier system.

L114 ANSWER 4 OF 15 HCPLUS COPYRIGHT 1999 ACS  
 AN 1998:135049 HCPLUS  
 DN 128:196638  
 TI Characteristics of hydrophobic-hydrophilic multiblock copolymers  
 AU Kim, Yong Joo; Sung, Yong Kiel; Piao, Ai Zhi; Kim, Sung Wan  
 CS Reliability Technology Division, NITQ, Kyungido, 427-010, S. Korea  
 SO Korea Polym. J. (1997), 5(4), 214-220  
 CODEN: KPJOE2; ISSN: 1225-5947  
 PB Polymer Society of Korea  
 DT Journal  
 LA English  
 AB ABCBA type amphiphilic block copolymers comprising poly(di-Me siloxane), poly(ethylene oxide), and heparin segments were synthesized by coupling reactions between intermediate and end-functionalized oligomers. These multiblock copolymers were identified using <sup>1</sup>H-NMR and FT-IR. The compns. of synthesized block copolymers were detd. by end group anal., sulfur elemental anal., and integration of NMR signals. The degrees of swelling of the multiblock copolymers increased by increasing the mol. wt. of PEO in water. By utilizing Wilhelmy plate contact angle measurement and electron microscopy, the surfaces of the block copolymers were characterized. Contact angle measurements demonstrated the hydrophilic effect of PEO and heparin on the hysteresis of the block copolymers and the heparinized block copolymers.

L114 ANSWER 5 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
AN 1997:592685 HCAPLUS  
DN 127:267968  
TI Evaluation of biological responses to polymeric biomaterials by RT-PCR analysis II: study of HSP 70 mRNA expression  
AU Kato, Shinya; Akagi, Takami; Kishida, Akio; Sugimura, Kazuhisa; Akashi, Mitsuru  
CS Dep. Applied Chemistry Chemical Engineering, Faculty Engineering, Kagoshima Univ., Kagoshima, 890, Japan  
SO J. Biomater. Sci., Polym. Ed. (1997), 8(10), 809-814  
CODEN: JBSEEA; ISSN: 0920-5063  
PB VSP  
DT Journal  
LA English  
AB In order to investigate how cells recognize biomaterials, mRNA that was expressed in attached HeLa S3 cells on various substrates was evaluated. As culture substrates, cellulose, ethylene-vinyl alc. copolymer (EVAL), nylon, tissue culture polystyrene (TCPS), high-d. polyethylene (PE), silicone rubber, and tetrafluoroethylene-hexafluoropropylene copolymer (yF) were used. HeLa S3 cells were cultured on these substrates for 24 h. The expressed HSP 70s mRNA was then isolated and detected using the RT-PCR method. As a result, the expression of HSP 70B mRNA was largely induced in cells that adhered to hydrophilic surfaces. On the other hand, on hydrophobic surfaces, the HSP 70B mRNA expression was low. It is concluded that HSP 70B mRNA expression is sensitive to differences in the hydrophilicity-hydrophobicity of the substrates.

L114 ANSWER 6 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
AN 1996:750669 HCAPLUS  
DN 126:135519  
TI Delivery of antitumor compounds to the rat colon: in vitro and in vivo evaluation  
AU Ciftci, Kadriye; Groves, Michael J.  
CS Institute for Tuberculosis Research, College of Pharmacy, University of Illinois at Chicago (M/C 964), 950 South Halsted Street, Room 2014 SEL, Chicago, IL, 60607-7019, USA  
SO Int. J. Pharm. (1996), 145(1,2), 157-164  
CODEN: IJPHDE; ISSN: 0378-5173  
PB Elsevier  
DT Journal  
LA English  
AB Using a rat model, the authors demonstrated that an enteric-coated hydroxypropyl Me cellulose (HPMC) granular formulation was capable of targeting or persisting in the colonic region. The formulation was optimized by measuring the in vitro release of 5-fluorouracil (5-FU) of granules prep'd. with different mol. wts. of HPMC ('Methocel') coated with different hydrophobicities of acrylic acid copolymers ('Eudragit'), a 'Methocel' K100M granule coated with 'Eudragit'-S being selected. X-ray examn. of lightly anesthetized rats demonstrated that orally administered enteric-coated granules contg. 50% wt./wt. barium sulfate persisted in the colon for longer than similar barium sulfate suspensions. Granules of HPMC, coated and uncoated, contg. 5-FU were administered by oral gavage and the tissue levels of drug were detd. by high performance liq. chromatog. At 6 h, drug from the uncoated formulation could be found in all tissues examd. On the other hand, at 8 h, drug from the coated granules could only be found in significant quantities in colon contents and colon tissue homogenates with increasing amts. being measured at 12 and 24 h. These data suggest that, at least in

the rat model, formulations can be designed that would persist in the colon and rectal regions, releasing drug to and not through the tissues. This concept might be valuable in the post-surgical treatment of colonic cancer, reducing the required dose of drug and therefore side effects. This should improve patient compliance and thus, the treatment outcome.

L114 ANSWER 7 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
 AN 1996:604906 HCAPLUS  
 DN 125:283803  
 TI Inhibitory effects of palmitic acid on anaerobic bacteria  
 AU Chu, Chun Feng; Miyahara, Takashi; Noike, Tatsuya  
 CS Tohoku Univ., Sendai, 980-77, Japan  
 SO Kogyo Yosui (1996), 455, 16-21  
 CODEN: KOYOAW; ISSN: 0454-1545  
 DT Journal  
 LA Japanese  
 AB Anaerobic decompn. of cellulose, glucose, gelatin, acetate, and propionate by inoculated wastewater was inhibited by the addn. of palmitate to the wastewater. Bacteria acclimatization was carried out for 3 mo to 2 yr according to the substrate property. Hydrophobicity of acclimatized sludge was measured by modified Matsumoto's method. Cellulose decompn. was strongly inhibited by palmitate, and inhibition decreased by the following order: glucose > acetate > propionate > gelatin. Sludge having weaker hydrophobicity was inhibited stronger by palmitate than the one having stronger hydrophobicity.  
 IT 57-10-3, Palmitic acid, processes  
 RL: ADV (Adverse effect, including toxicity); BUU (Biological use, unclassified); PEP (Physical, engineering or chemical process); PRP (Properties); BIOL (Biological study); PROC (Process); USES (Uses) (inhibitory effects of palmitic acid on org. substrate biodegrdn. in anaerobic wastewater treatment)

L114 ANSWER 8 OF 15 HCAPLUS COPYRIGHT 1999 ACS  
 AN 1996:576363 HCAPLUS  
 DN 125:256905  
 TI Chemical modifications of heparin. II. Hydrophobization of partially N-desulfated heparin  
 AU Diancourt, Francis; Braud, Christian; Vert, Michel  
 CS C.R.B.A., Faculte de Pharmacie, Montpellier, 34060, Fr.  
 SO J. Bioact. Compat. Polym. (1996), 11(3), 203-218  
 CODEN: JBCPEV; ISSN: 0883-9115  
 DT Journal  
 LA English  
 AB In order to hydrophobize heparin, dodecanal, cholic acid and stearic acid were conjugated, resp., to partially N-desulfated heparin with the formation of imine bonds in the case of dodecanal and amide bonds in the case of the two latter compds. It was found that the three conjugates were aggregated in aq. soln. with the formation of .apprx.300 nm aggregates. The conjugates retained the anticoagulant activity of the parent heparin according to in vitro tests. In vivo, slightly prolonged activity was obsd. after s.c. injection. The activity lasted for at least 2 h after i.v. injection suggesting that aggregates of hydrophobized heparin were not captured by the reticuloendothelial system. These new water dispersed aggregates are of potential interest for the transport and slow release of hydrophobic drugs in blood either as macromol. prodrugs or after mol. microencapsulation, and for oral administration.  
 IT 57-11-4DP, Stearic acid, conjugates with

**desulfated heparin 81-25-4DP, Cholic acid, conjugates with desulfated heparin**  
**RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process)**  
 (hydrophobization of partially desulfated heparin in relation to anticoagulant activity)

IT 57-11-4, Stearic acid, reactions

81-25-4, Cholic acid

RL: RCT (Reactant)

(hydrophobization of partially desulfated heparin in relation to anticoagulant activity)

L114 ANSWER 9 OF 15 HCAPLUS COPYRIGHT 1999 ACS

AN 1996:113950 HCAPLUS

DN 124:211942

TI Blood compatible biomaterials: hydrophilicity vs. hydrophobicity

AU Sevastianov, Viktor I.; Drushlyak, Irina V.; Eberhart, Robert C.; Kim, Sung Wan

CS Institute Transplantology Artificial Organs, Moscow, Russia

SO Macromol. Symp. (1996), 103(Polymers and Medicine), 1-4

CODEN: MSYMEC; ISSN: 1022-1360

DT Journal

LA English

AB A tendency towards unification of adsorption properties of hemo compatible hydrophilic and hydrophobic surfaces within the first minutes of contact with protein was demonstrated. Nevertheless, the change of adsorption properties of blood compatible hydrophilic surfaces after passivation with proteins is much less in comparison with the one for blood compatible hydrophobic surface.

L114 ANSWER 10 OF 15 HCAPLUS COPYRIGHT 1999 ACS

AN 1995:731156 HCAPLUS

DN 123:122944

TI Applications of hydrophobically-modified hydroxypropyl methyl cellulose to topical preparations, IV. Influence of indomethacin and isopropanol on hydrophobic interaction through long-chain alkyl groups of hydrophobically-modified hydroxypropyl methyl cellulose

AU Ikeda, Kaori; Saitoh, Izumi; Oguma, Takayoshi; Takagishi, Yasushi

CS Developmental Res. Labs., Shionogi and Co., Ltd., Hyogo, 660, Japan

SO Chem. Pharm. Bull. (1995), 43(6), 1012-16

CODEN: CPBTAL; ISSN: 0009-2363

DT Journal

LA English

AB The hydrophobic interaction between long-chain alkyl groups of hydrophobically-modified hydroxypropyl Me cellulose (HM-HPMC) and indomethacin (IM) was studied using a fluorescence probe method in which pyrene mols. are allowed to assoc. with long-chain alkyl groups. In an HM-HPMC soln., the hydrophobicity of the pyrene environment markedly increased as the concn. of HM-HPMC was increased, indicating an increase of hydrophobic interaction between the long-chain alkyl groups. When IM was added to this soln., the hydrophobicity of the pyrene environment decreased as the IM concn. increased. Thus, it was suggested that the hydrophobic interaction between long-chain alkyl groups and IM caused a weakening of the hydrophobic interaction between HM-HPMC mols. themselves. When isopropanol (IPA) was added to an HM-HPMC soln., IPA weakened the hydrophobic interaction through long-chain alkyl groups. Thus, IPA caused an expansion of the intermol. hydrophobic regions. For this reason, the amt. of IM entrapped in these regions increased when the IPA concn. as

increased. However, when the IPA concn. was further increased, the hydrophobic interaction finally disappeared. Thus, an IM mols. could no longer be trapped in the hydrophobic regions, the activity of HM-HPMC suppressing the growth of IM crystals disappeared.

L114 ANSWER 11 OF 15 HCPLUS COPYRIGHT 1999 ACS  
AN 1994:426516 HCPLUS  
DN 121:26516  
TI New Approaches for the Preparation of Hydrophobic Heparin Derivatives  
AU Liu, Jian; Pervin, Azra; Gallo, Cindy M.; Desai, Umesh R.; Van Gorp, Cornelius L.; Linhardt, Robert J.  
CS College of Pharmacy, University of Iowa, Iowa City, IA, 52242, USA  
SO J. Pharm. Sci. (1994), 83(7), 1034-9  
CODEN: JPMSAE; ISSN: 0022-3549  
DT Journal  
LA English  
AB A **heparin** deriv. sufficiently lipophilic to be bound to plastics, forming blood-compatible supports, or to be used as an anticoagulant by transdermal or oral routes would be of great pharmaceutical interest. For such applications, the functional groups within **heparin**'s antithrombin III binding site, responsible for its anticoagulant activity, cannot be modified. Chem. is described in which lipophilic substituents were attached to the reducing termini of **heparin** chains. Substituents introduced at this position had a minimal effect on the antithrombin III binding sites found in **heparin**'s interior. These derivs., with enhanced hydrophobicities, were prepd. using two distinctly different approaches. First, octyl isocyanate and octadecyl isocyanate were coupled to the core peptide of peptidoglycan **heparin** to form octyl- and octadecyl-peptidoglycan **heparin**. These octyl- and octadecyl-peptidoglycan **heparins** were then purified by hydrophobic interaction chromatog. on phenyl-Sepharose CL-4B, demonstrating their enhanced hydrophobicities. Second, the lipophilic acyl hydrazides of various long chain fatty acids were coupled to **heparin**'s reducing end. Caprylic (C8), capric (C10), lauric (C12), and stearic(C18) hydrazide derivs. of **heparin** were prepd. using this approach. Only the stearyl hydrazide deriv. of **heparin** showed a measurable increase in lipophilicity. This result demonstrated that a single small linear C8, C10, or C12 aliph. chain was ineffective in enhancing the hydrophobicity of the highly neg., polyanionic **heparin** mol. Two lipophilic chains, lauryl (C12) and stearyl (C18), were then coupled to a single **heparin** chain, resulting in a **heparin** deriv. having enhanced hydrophobicity. All the **heparin** derivs. prepd. in this study maintained some of their anticoagulant activity.

L114 ANSWER 12 OF 15 HCPLUS COPYRIGHT 1999 ACS  
AN 1993:415249 HCPLUS  
DN 119:15249  
TI In vitro passive and iontophoretically assisted transport of salbutamol sulfate across synthetic membranes  
AU Rodriguez Bayon, A. M.; Corish, J.; Corrigan, O. I.  
CS Fac. Farm., Univ. Complutense Madrid, Madrid, 28040, Spain  
SO Drug Dev. Ind. Pharm. (1993), 19(10), 1169-81  
CODEN: DDIPD8; ISSN: 0363-9045  
DT Journal  
LA English  
AB The passive and elec. assisted transport of salbutamol sulfate through

four synthetic membranes was investigated. Two of these were hydrophilic (Visking 18/32 and Celgard-3401) and two hydrophobic (Celgard-2400 and Celgard-4500). Significant differences in passive membrane transport were obsd. The hydrophilic Celgard membrane gave similar passive transport rates to Visking 18/32. However, slower rates were obsd. with the hydrophobic membranes, the rate for Celgard-4500 being 4-5 fold smaller than Visking 18/32 and that for Celgard-2400 being negligible over a period of 6 h. The passive release of salbutamol sulfate from the hydrogel across the hydrophilic membranes was matrix-controlled, whereas the membrane was the rate-limiting element for passive release through the hydrophobic membranes. Application of an elec. potential giving rise to iontophoretic currents in the range 0.100 to 0.500 mA led to an increase in drug transport rate and this effect became larger as the current was increased. The quantity of drug transported in a given time period increased linearly with time for both kind of membranes. However, the relative increase in transport induced by current was greatest with the hydrophobic membrane.

- L114 ANSWER 13 OF 15 HCPLUS COPYRIGHT 1999 ACS  
AN 1991:12177 HCPLUS  
DN 114:12177  
TI Protein adsorption from semi-permeable membranes during biologic solution concentration through effective dialysis  
AU Godevirogova, Ts.; Dimov, A.; Petrov, S.  
CS Dep. Gen. Chem. Technol., Higher Chem. Tech. Inst., Burgas, 8010, Bulg.  
SO Dokl. Bolg. Akad. Nauk (1990), 43(8), 149-52  
CODEN: DBANAD; ISSN: 0366-8681  
DT Journal  
LA English  
AB Adsorption isotherms of membranes against human albumin having concns. 0-12 g/L during 12 h adsorption under static dialysis conditions without using cellulose filters showed that the highest amt. of proteins were adsorbed by polysulfone membranes which had the highest hydrophobicity while the lowest amt. of proteins were adsorbed by regenerated cellulose and partially hydrolyzed polyacrylonitrile membranes.
- L114 ANSWER 14 OF 15 HCPLUS COPYRIGHT 1999 ACS  
AN 1990:558657 HCPLUS  
DN 113:158657  
TI Examination of the permeability of microfiltration membranes  
AU Moiseenko, L. A.  
CS VNII Antibiot, Moscow, USSR  
SO Khim.-Farm. Zh. (1990), 24(6), 64-7  
CODEN: KHFZAN; ISSN: 0023-1134  
DT Journal  
LA Russian  
AB The permeability of microfiltration membranes depended on the nature of a polymer method of producing a porous film, and hydrophobicity. Wettability affected the diffusion of liqs. through the hydrophobic membranes. An equation for calcg. the membrane permeability is given.

- L114 ANSWER 15 OF 15 HCPLUS COPYRIGHT 1999 ACS  
AN 1989:601474 HCPLUS  
DN 111:201474  
TI Effects of the hydrophobicity of glass beads surface and moisture content of binder on the hardness of tablets  
AU Wada, Yasutaka; Mizuta, Taiiti; Sakamoto, Teruo  
CS Res. Lab., Shionogi Co., and Ltd., Ohsaka, 553, Japan

SO Yakugaku Zasshi (1989), 109(7), 480-6  
CODEN: YKKZAJ; ISSN: 0031-6903

DT Journal  
LA Japanese

AB Tablet hardness was measured for direct compressed tablets contg. various glass beads and binders. Intact, trimethylsilylated and chloromethyldimethylsilylated glass beads were used to investigate the effects of surface property on the tablet hardness. The tablet hardness was decreased by the hydrophobic modification of glass beads surface. The increase of moisture contents of binder accelerated the decrease of tablet hardness of trimethylsilylated glass beads tablets. From SEM, it was concluded that the decrease of tablet hardness obsd. for the hydrophobic glass beads could be explained in terms of the variation of the adhesive ability of binder particles to glass beads.